

Case Report

Breast cancer metastasising to the uterine cervix

C A Bryson, R H B de Courcy-Wheeler, R J Wallace

Accepted 28 January 1999

CASE REPORT A 47-year-old mother of two was seen in the colposcopy clinic in November 1994 with a history of two abnormal smears. The first in June 1993 showed severe inflammation, and the repeat smear a year later showed atypical glandular cells. She had been post-menopausal for two years and had no gynaecological symptoms. In June 1993, a left mastectomy with axillary node sampling had been performed for a well-differentiated, infiltrating ductal carcinoma of the breast (T1 N0 M0). There was extension to the pectoralis major muscle but no lymph node involvement. She had been on Tamoxifen 20 mg daily since the operation.

At colposcopy the vagina was atrophic, and the epithelium was sloughing. The cervix felt hard and irregular. There was a large iodine-negative area but no evidence of cervical intraepithelial neoplasia. She was admitted for examination under anaesthesia and cervical biopsy. The uterus and adnexae were unremarkable and no induration could be felt beyond the cervix. A 1 x 0.5 cm biopsy showed extensive involvement of the cervix with well-differentiated adenocarcinoma and lymphatic permeation. A computerised tomography scan of the chest, abdomen and pelvis was normal as were a bone scan, magnetic resonance imaging of the pelvis and liver function tests.

A total abdominal hysterectomy and bilateral salpingo-oophorectomy were performed in January 1995. Histology confirmed metastatic breast carcinoma confined to the cervix. Post operative recovery was complicated by a lower respiratory tract infection which responded to antibiotics. She received 22 fractions of radiotherapy to her anterior, posterior, right lateral and left lateral pelvis (4400 centigray), followed by six courses of chemotherapy consisting of 5-fluorouracil, epirubicin and cyclophosphamide. She remains well to date almost 4 years after operation.

DISCUSSION

The cervix is not usually suspected of being a potential site for metastatic disease, is less likely to be examined at post-mortem, and adenocarcinoma metastatic to the uterine cervix is rare. The reasons for its rarity are that the cervix is a small target organ with a small blood supply and only an afferent lymph drainage system. Further, the fibromuscular tissue of the cervix is a poor culture medium. The response of the cervix to metastatic disease is fibrous proliferation and an inflammatory cellular reaction which may explain the findings in this case of an expanded, indurated cervix.

Most extragenital metastases to the cervix arise from primary tumours of the gastrointestinal tract.¹ Other primary sites include lung, pancreas, melanoma, urethra and breast. Breast cancer often metastasises to the ovary but rarely to the cervix. It has been estimated that only 4.5% of cervical secondaries arise from breast primaries.²

Breast cancer is one of the most common tumours affecting women, representing about 30% of all female malignancies. In 1941, 34 of 59 (58%) cases of carcinoma metastatic to the uterus and cervix originated in the breast.³ Regrettably these

Department of Obstetrics and Gynaecology, Craigavon Area Hospital, 68 Lurgan Road, Portadown BT63 5QQ.

Caroline Bryson, MRCOG, Specialist Registrar in Obstetrics and Gynaecology.

R J Wallace, FRCOG, Consultant Obstetrician Gynaecologist.

Department of Obstetrics and Gynaecology, Daisy Hill Hospital, Newry BT35 8DR.

R H B de Courcy-Wheeler, MD, MRCOG, Consultant Obstetrician Gynaecologist.

Correspondence to Dr Bryson.

TABLE

Details of 27 cases of breast cancer metastasising to the uterine cervix.

<i>Case No.</i>	<i>Reference</i>	<i>Age</i>	<i>Interval months</i>	<i>Symptoms</i>	<i>Survival months</i>
1	9 Cruz 1954	47	60	menorrhagia	alive
2	10 Wallach & Edberg 1957	73	6	facial palsy	0.5
3	11 Song 1963	45	0	smear + IMB	alive
4	„	49	0	IMB	alive
5	„	51	48	smear+ IMB	4
6	12 Dance & Fullmer 1970	63	48+	smear + nodules	alive
7	13 Cohan & Kaplan 1984	74	12	PMB	alive
8	14 Di Bonito et al 1985				
9	„	–	108	–	108
10	„	–	–	PVB	?
11	„	–	12	PVB	12
12	„	–	5	PVB	5
13	„	–	36	none	36
14	5 Lemoine & Hall 1986	45	36	PMB	1.5
15	„	60	58	PMB	4
16	„	39	-2	IMB	6
17	„	71	58	PMB	14
18	15 Way 1980	49	2	PVB	155
19	16 Korhonen & Stenback 1984	63	8	smear + PMB	alive
20	6 Yazigi et al 1988	49	0	PVB	14
21	„	49	42	PVB	2
22	„	38	0	breast lump	15
23	17 Taylor & Torode 1990	84	444	PMB	1
24	„	50	33	IMB + polyp	alive
25	18 Campora et al 1991	57	83	abdominal pain	6
26	4 Fiorella et al 1992	54	0	smear + PMB	6
27	Current case	47	17	smear	alive

Interval = interval between diagnosis of primary breast cancer and secondary cervical cancer

IMB = intermenstrual bleeding. PMB=post-menopausal bleeding. PVB=peri-vaginal bleeding.

Alive = alive at time of report

cases were poorly documented, and no differentiation was made between the cervix and the body of the uterus. Since then a further twenty-seven cases of breast carcinoma metastasising to the cervix (including this one) have been reported in the literature. Age at presentation, interval between primary and secondary diagnosis, presenting symptoms and survival are summarised in the Table.

Most cases presented with abnormal vaginal bleeding. Seven cases had abnormal cervical cytology but only the current case presented solely with an abnormal smear. Atypical glandular cells in a cervical smear suggest endometrial carcinoma, but they may represent any adenocarcinoma primary. The cytological appearances, in particular the absence of a tumour diathesis, may indicate an extragenital site.⁴ In one case the cervical metastasis was detected two months before the breast cancer could be identified. However, the average interval between primary and secondary diagnosis was 44.5 months. In comparison to other primaries the relatively long interval for breast disease has already been highlighted.⁵

In most reports there were multiple metastases. In this case and in one other report⁶ the cervix was the only site of secondary disease. On initial presentation it is easy to mistake cervical metastases as primary disease because of the common presentation of abnormal vaginal bleeding, abnormal cervical smear and an abnormal-looking cervix. The histological diagnosis of a secondary deposit makes an important difference to treatment. This patient had adjuvant radiotherapy and chemotherapy which would not have been usual management for a primary adenocarcinoma of the cervix.

Had the cervix had been normal, the atypical glandular cells in the smear might have indicated an adenocarcinoma further up the genital tract. The annual incidence of endometrial carcinoma at ages 55-84 in Europe is 0.5 per 1000, which is doubled after two years of tamoxifen medication and approximately quadrupled after five years of tamoxifen.⁷ Abnormal vaginal bleeding in women taking tamoxifen needs rigorous investigation but the benefits of screening asymptomatic women for endometrial carcinoma with transvaginal ultrasound or hysteroscopy have yet to be determined.⁸

We suggest that any woman with a history of breast carcinoma who presents with gynaecological symptoms should be screened both for the iatrogenic effects of tamoxifen therapy on the endometrium and for metastatic disease of the cervix. Investigation of extragenital sites is particularly relevant if adenocarcinoma of the cervix is suspected.

REFERENCES

1. Mazur M T, Hsueh S, Gersell D J. Metastases to the female genital tract. analysis of 325 cases. *Cancer* 1984; **53**: 1978-84.
2. Ng A B, Teeple D, Lindner E A, Reagan J. The cellular manifestations of extrauterine cancer. *Acta Cytol* 1974; **18**: 108-17.
3. Charache H. Metastatic carcinoma in uterus. *Amer J Surg* 1941; **53**: 152-7.
4. Fiorella R V, Beckwith L G, Miller L K, Kragel P J. Metastatic signet ring carcinoma of the breast as a source of positive cervicovaginal cytology. a case report. *Acta Cytol* 1992; **37**: 948-52.
5. Lemoine N R, Hall P A. Epithelial tumours metastatic to the uterine cervix. *Cancer* 1986; **57**: 2002-5.
6. Yazigi R, Sandstad J, Munoz A K. Breast cancer metastasing to the uterine cervix. *Cancer* 1988; **61**: 2558-60.
7. Early Breast Cancer Trialists' Collaborative Group. Tamoxifen for early breast cancer: an overview of the randomised trials. *Lancet* 1998; **351**: 1451-67.
8. Bissett D, Davis J A, George W D. Gynaecological monitoring during tamoxifen therapy. *Lancet* 1994; **344**: 1244.
9. Cruz P. Metastatic carcinoma with metastasis to the cervix uteri. *Delaware Med J* 1954; **26**: 304-5.
10. Wallach J B, Edberg S. Carcinoma metastatic to the uterine cervix. *Am J Obstet Gynecol* 1959; **77**: 990-5.
11. Song J. Metastatic carcinoma of the uterine cervix from primary breast cancer. *JAMA* 1963; **184**: 498-500.
12. Dance E F, Fullmer C D. Extrauterine carcinoma cells observed in cervico-vaginal smears. *Acta Cytologica* 1970; **14**: 187-91.
13. Cohan I, Kaplan A L. Postmenopausal bleeding secondary to metastatic disease in the endocervix from carcinoma of the breast. *Gynaecol Oncol* 1984; **17**: 133-6.
14. Di Bonito L, Patriarca S, Alberico S. Breast carcinoma metastasizing to the uterus. *Eur J Gynaecol* 1985; **6**: 211-7.
15. Way S. Carcinoma metastatic in the cervix. *Gynecol Oncolo* 1980; **9**: 298-302.
16. Korhonen M, Stenbäck F. Adenocarcinoma metastatic to the uterine cervix. *Gynecol Obstet Invest* 1984; **17**: 57-65.
17. Taylor W G, Torode H W. Metastatic breast carcinoma in the pelvis. *Br J Clin Pract* 1990; **44**: 205-6.
18. Campora E, Serra G E, Nicolo G, Gatteschi B, Rosso R. Endocervical metastases secondary to breast carcinoma: a Case Report. *Eur J Gynaecol Oncol* 1991; **12**: 103-6.